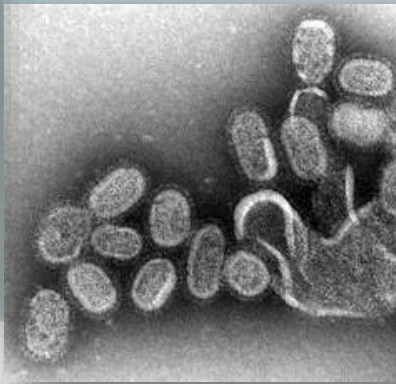


Update on Avian Influenza



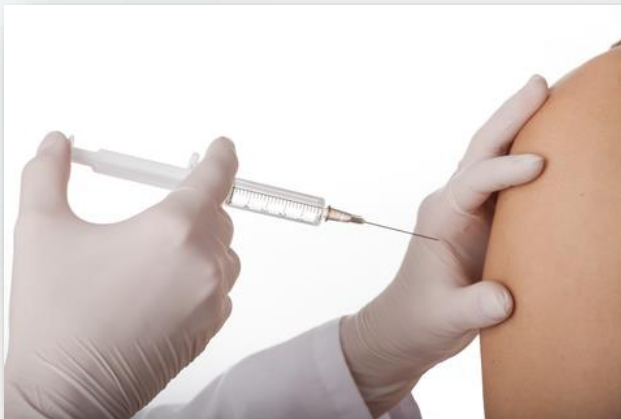
Mary J. Pantin-Jackwood, DVM, MS, PhD
Southeast Poultry Research Laboratory
U.S. Department of Agriculture, Athens, Georgia
mary.pantin-jackwood@ars.usda.gov

Overview

- Introduction: Differences between seasonal flu, pandemic flu and avian flu
- Influenza A viruses
- Avian influenza – Epidemiology and pathobiology (LPAI and HPAI)
- Avian influenza in commercial poultry – Current situation
- Asian lineage H5N1 HPAI
- H7N9 influenza in China
- Prevention and Control



Seasonal Influenza



Influenza virus type A and B

<http://www.cdc.gov/flu/index.htm>

Pandemic influenza



Emergency hospital, Camp Funston,
Kansas 1918

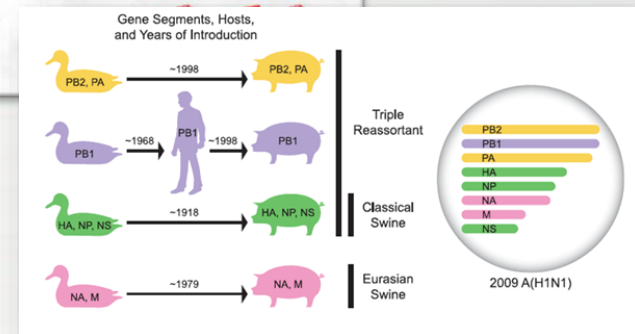
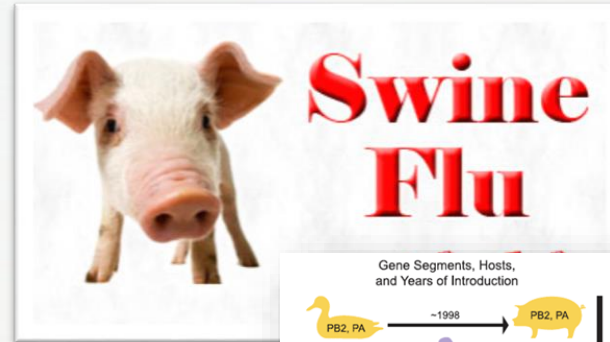
Courtesy of National Museum of Health and Medicine

1918 “Spanish Flu” (H1N1) 20-40 million deaths

1957 “Asian Flu” (H2N2) 1 million deaths

1968 “Hong Kong Flu” (H3N2) 1 million deaths

pH1N1

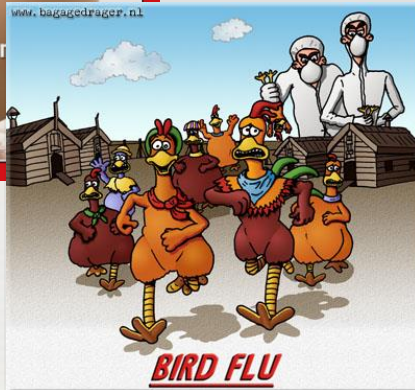


Nancy Cox, 2009, Science

Pandemic threats

H5N1 HPAI

668 total cases/393 deaths



H7N9 LPAI

464 total cases/174 deaths



China's deadly new H7N9 bird flu virus may be harder to track than predecessors, scientists say
AP/ April 3, 2013

H10N8 LPAI

3 total cases/2 deaths

Influenza in animals

H7N3 HPAI

Mexico Kills 8 Million Chickens to Contain H7N3 Virus

By C. Hsu | Aug 8, 2012 EDT



H3N2v

Health officials warn fair attendees not to pet pigs due to swine flu fears

Published August 31, 2012. FoxNews.com

H5N8 HPAI

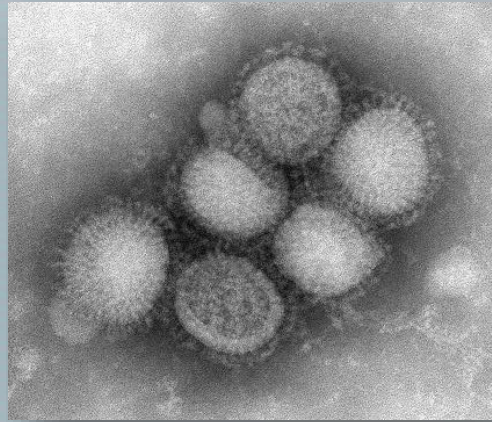
Bird flu worries spread in Europe

By S. Capelouto, CNN. November 24, 2014

H5N2 HPAI

Canada bird flu virus identified as 'highly pathogenic' strain.

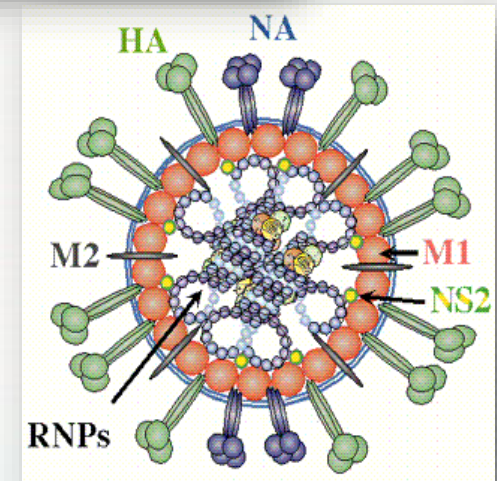
Published Dec. 5 2014 FoxNews.co



Influenza A viruses

Influenza A viruses

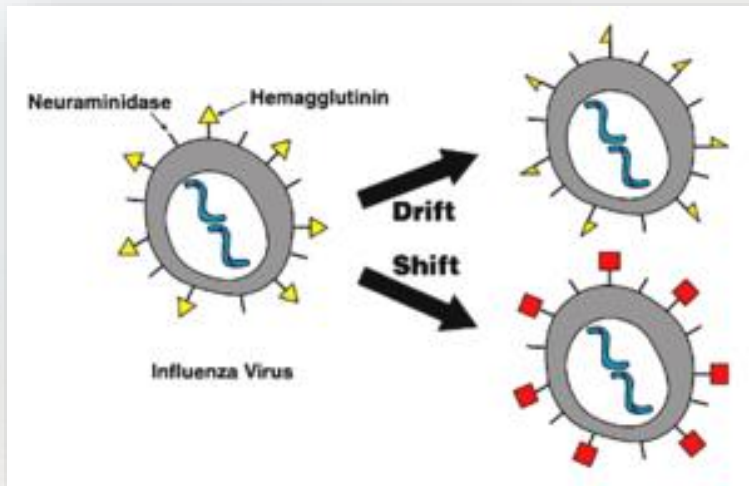
- Influenza A viruses infect humans, birds and other animals including horses, pigs, dogs
- The natural reservoir of influenza A viruses is wild aquatic birds
- Classified based on the viral surface proteins hemagglutinin (HA or H) and neuraminidase (NA or N)
 - 16 H subtypes (or serotypes) and 9 N subtypes



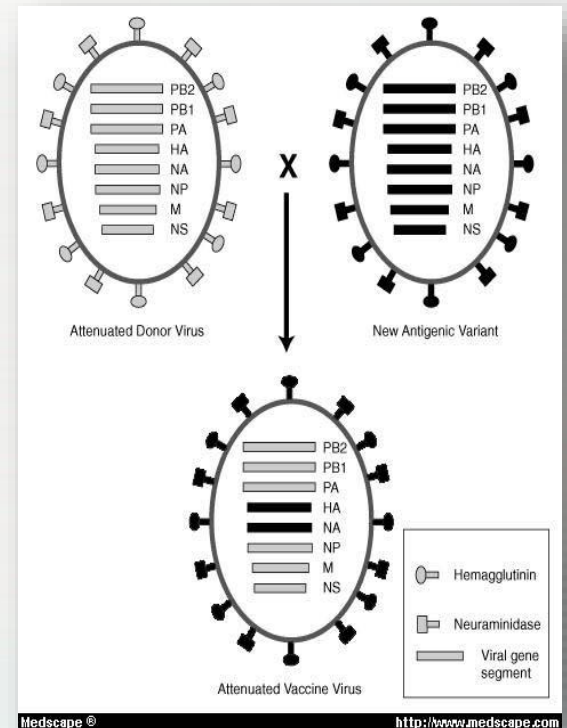
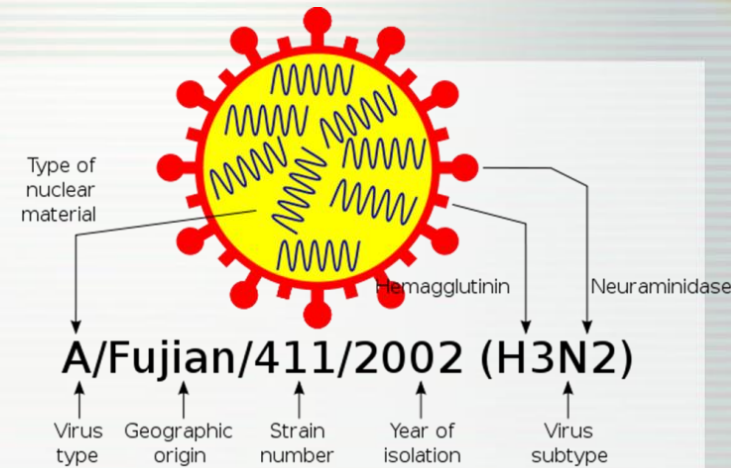
© Paul Digard, Dept Pathology,
University of Cambridge.

Influenza A viruses

- Orthomyxoviridae family, genus Influenzavirus type A
- Single-stranded (-) sense segmented RNA genome



New influenza A viruses are constantly emerging



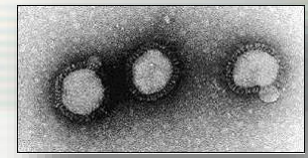
Reassortment

Influenza A viruses

Wide host range, strain dependent



Influenza A viruses

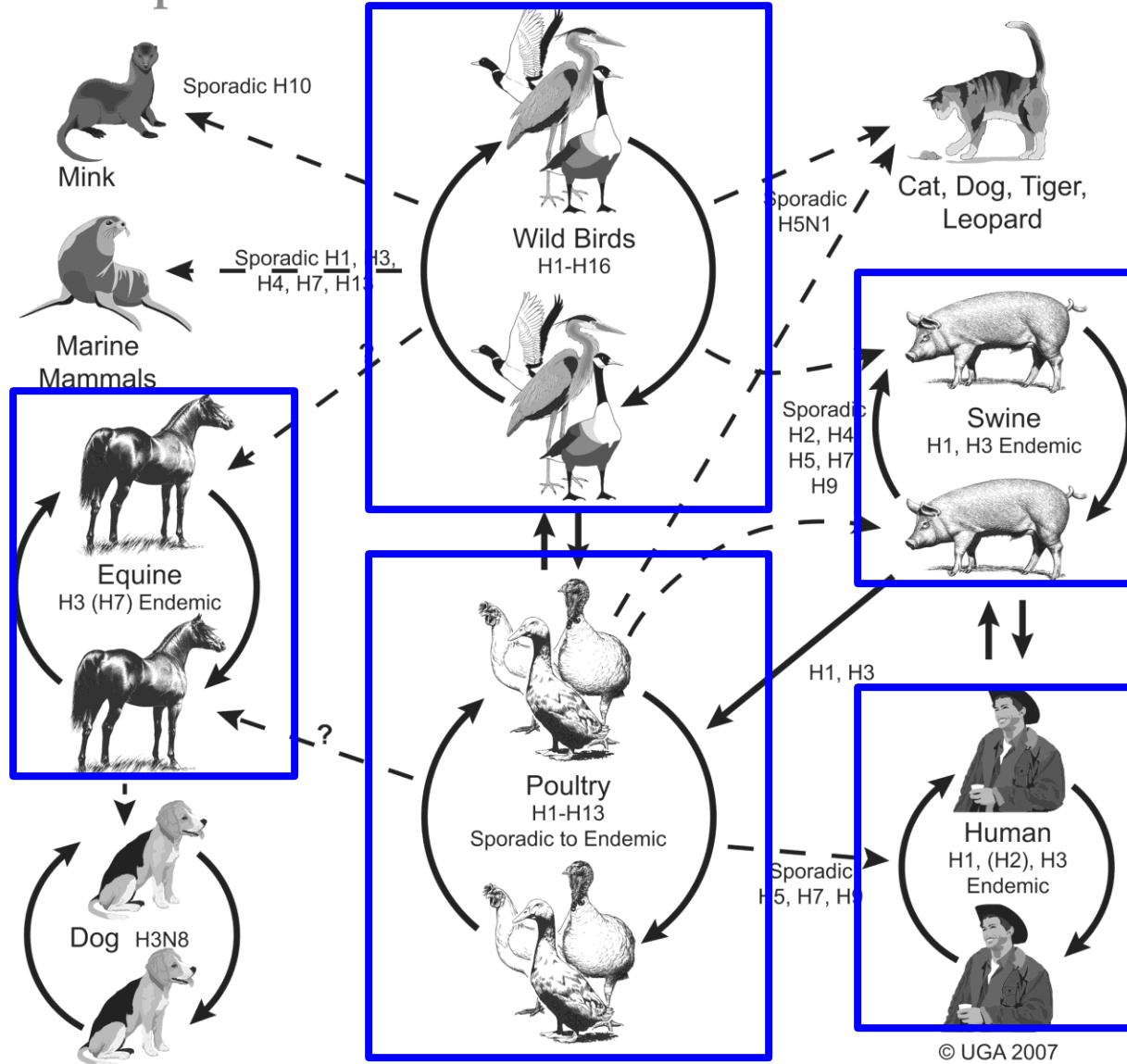


Subtype	Mammalia			Aves		
	Human	Swine	Equine	Wild Ducks	Shorebirds	Poultry
H1	+	++		+	+	++
H2	(+)			+	+	+
H3	+++	++	++	++		+
H4		±		++	+	+
H5	±	±		+	+	++
H6				++	+	+
H7	±		(+)	+	+	++
H8				+		+
H9	±	±		+	++	++
H10				+	+	+
H11				+	+	+
H12				+	+	
H13					++	+
H14				+		
H15				+	+	
H16					+	

H17 – type A influenza from bats

Adapted from Swayne, D.E. Epidemiology of Avian Influenza in Agricultural and Other Man-Made Systems. In: Avian Influenza. Wylie-Blackwell, March, 2008.

Interspecies Transmission of Influenza A Viruses



Swayne, D.E. Epidemiology of Avian Influenza in Agricultural and Other Man-Made Systems. In: Avian Influenza. Wylie-Blackwell, March, 2008.



Avian Influenza

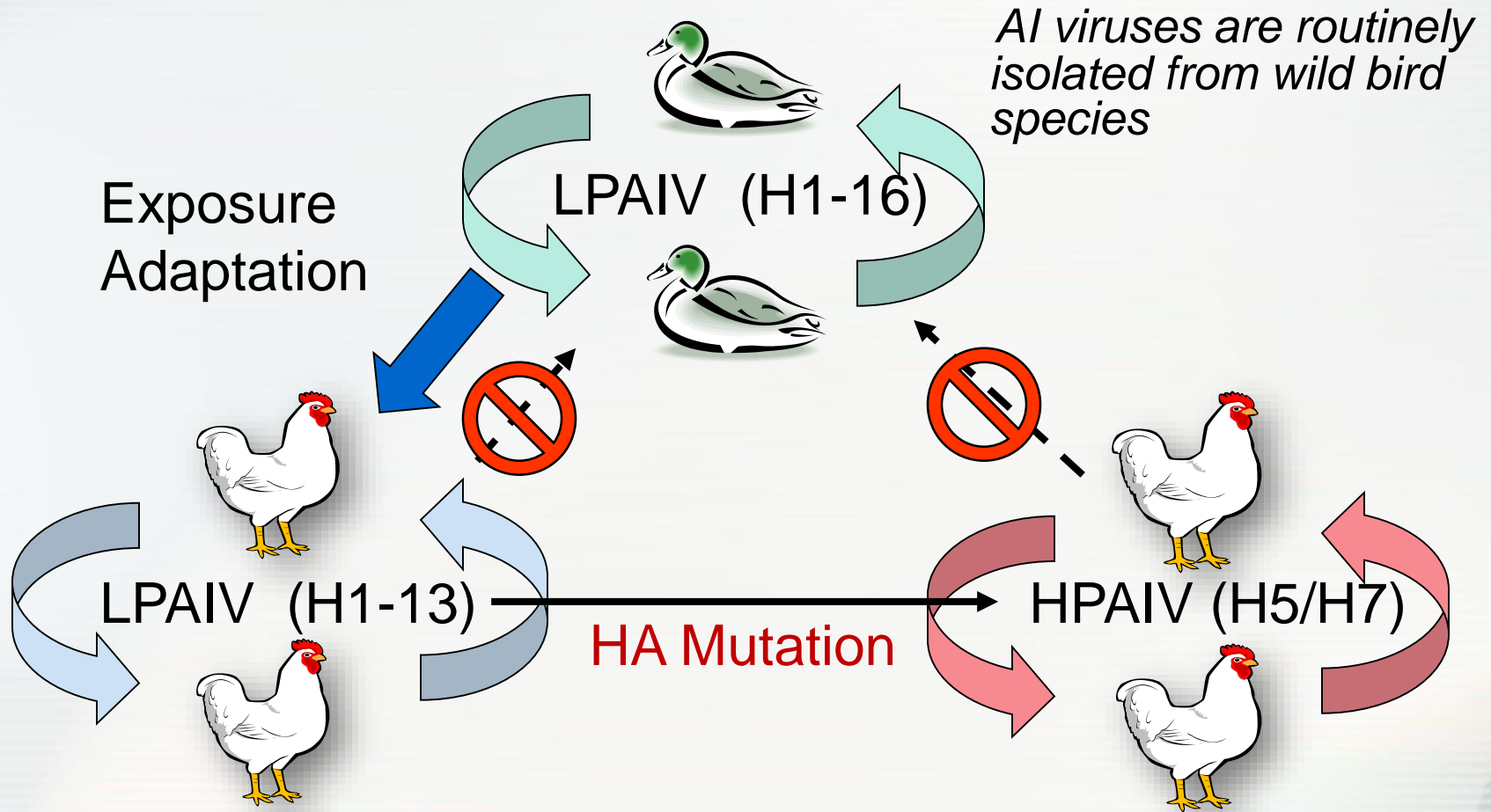
Avian influenza virus: Pathogenicity

- Pathogenicity: Ability to produce lesions, disease and/or death in a host
- Determined by several factors: virus, host, and environmental
- Host differences – *Galliformes* vs. *Anseriformes*
- Pathotypes:
 - High pathogenicity (HPAI): some H5 and H7 viruses
 - Low pathogenicity (LPAI): most H1-13 virus

LPAI H5 or H7 subtypes can mutate into HPAI viruses



AIV: Ecology and Epidemiology



The vast majority of HPAI viruses never re-enter the wild bird system

What do we mean by highly pathogenic?

Only applies to gallinaceous poultry (chickens, turkeys, quail)

– Regulatory definitions from OIE

1. Highly pathogenic avian influenza (HPAI) virus lethal for 75% or more of intravenously inoculated susceptible chickens (IVPI > 1.2) within 10 days.
2. H5 or H7 subtype that has an amino acid sequence at the hemagglutinin cleavage site 'compatible' with HPAI due to multiple basic amino acids

- Notifiable avian influenza
 - H5/H7 HPAI
 - H5/H7 low pathogenicity notifiable AI (LPNAI)
- Other low pathogenicity avian influenza viruses are not notifiable (H1-4, H6, H8-16)



World Organization for Animal Health

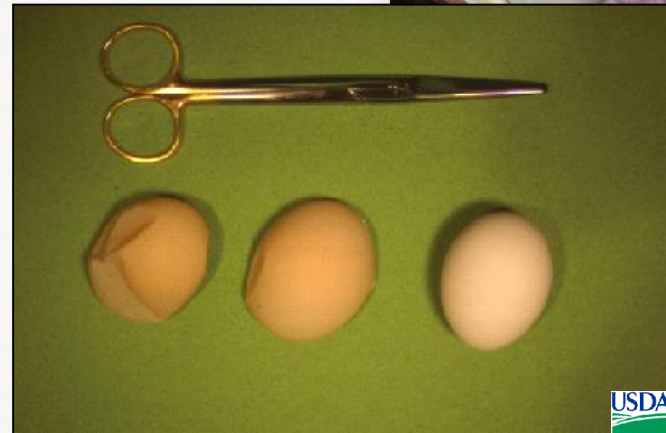


Low Pathogenicity Avian Influenza

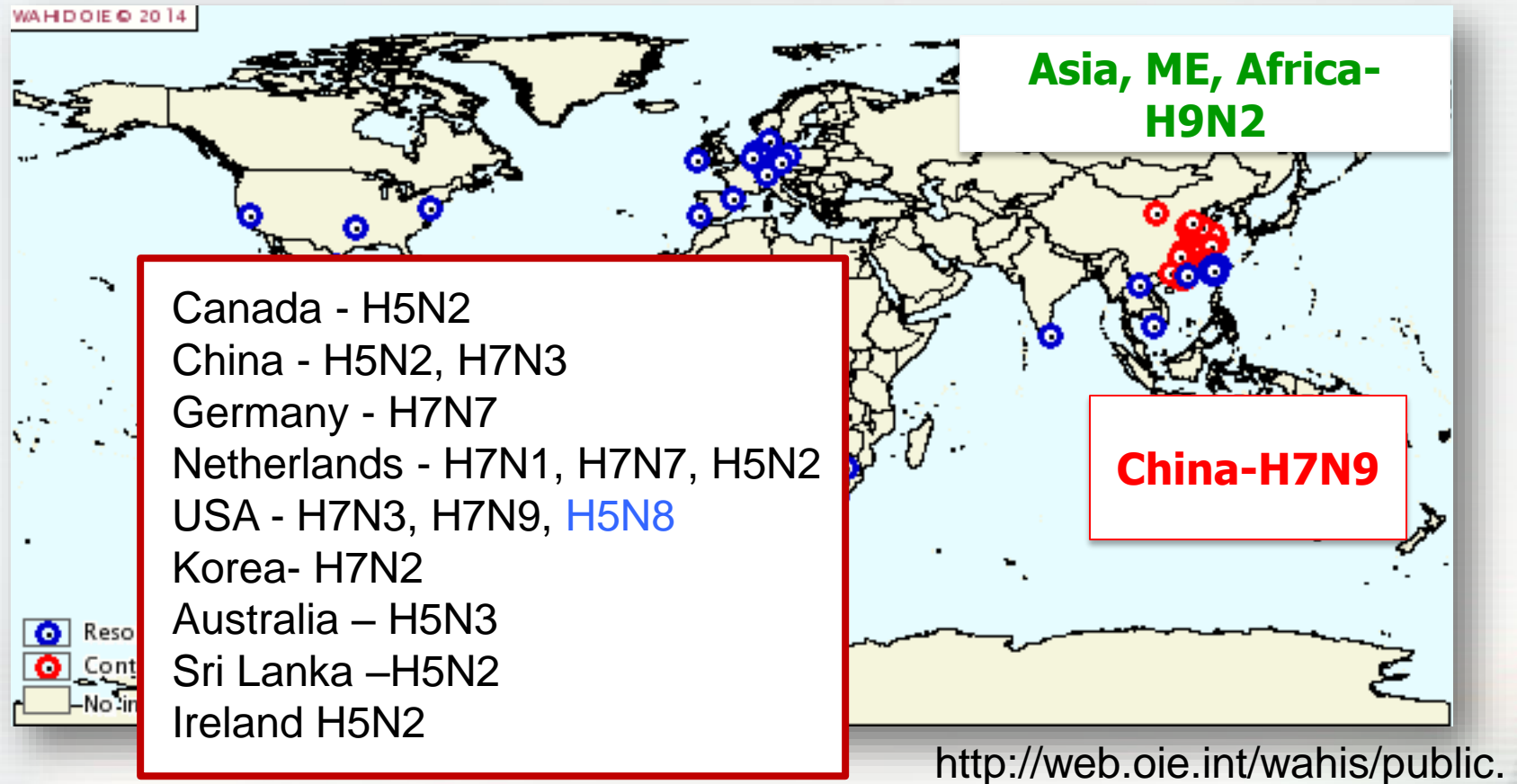
Chickens and turkeys

Infections - variable clinically

- Respiratory signs
- Rhinitis and tracheitis
 - Decrease in feed and water consumption
- Ocular discharge
- Diarrhea
- Drops in egg production
- Mild increase in mortality



LPAI outbreaks reported to the OIE 2012-2014



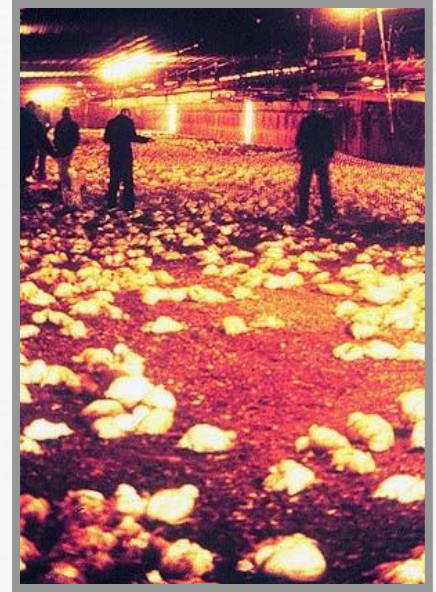
H5 and H7 avian influenza in its low pathogenic form in poultry is a notifiable disease as per Chapter 10.4. on avian influenza of the Terrestrial Animal Health Code (2010)

Highly Pathogenic Avian Influenza

Chickens and turkeys

Severe systemic disease. Highly contagious

- High mortality. Rapid spread
- Severe decrease in feed and water consumption
- +/- nervous signs
- Edema of head and legs
- Pulmonary edema, congestion and hemorrhage
- Visceral hemorrhage



37 HPAI Disease Events

1890' s-1950' s numerous outbreaks, subtypes not known
1924-25, North east US H7N7?

1. 1959-Scotland, H5N1

2. 1961-S. Africa, H5N3

3. 1963-England, H7N3

4. 1966-Canada, H5N9

5. 1975-Australia, H7N7

6. 1979 –Germany, H7N7

7. 1979-England, H7N7

***8. 1983-84 - USA, H5N2**

9. 1983-Ireland, H5N8

10. 1985-Australia, H7N7

11. 1991-England, H5N1

12. 1992-Australia, H7N3

13. 1994-Australia, H7N3

***14. 1994-95-Mexico, H5N2**

15. 1995 **and 2004** –Pakistan, H7N3

16. 1997-Australia, H7N4

17. 1997-Italy, H5N2

***LPAIV ⇒HPAIV**

****Largest epizootic on record**

****18. 1996-present – Asia/Europe/
Africa, H5N1**

***19. 1999-2000 - Italy, H7N1**

***20. 2002 - Chile, H7N3**

***21. 2003 – Netherlands, H7N7**

***22. 2004 – USA, H5N2**

***23. 2004 – Canada, H7N3**

24. 2004, 2006 – S. Africa, H5N2

25. 2005? – N. Korea, H7N7

***26. 2007 – Canada, H7N3**

27. 2008 – England, H7N7

28. 2009- Spain, H7N7

29. 2011- South, Africa H5N2

30. 2012- Taipei, H5N2

31. 2012- South Africa, H7N1

32. 2012. Mexico, H7N3

33. 2012: Australia, H7N7

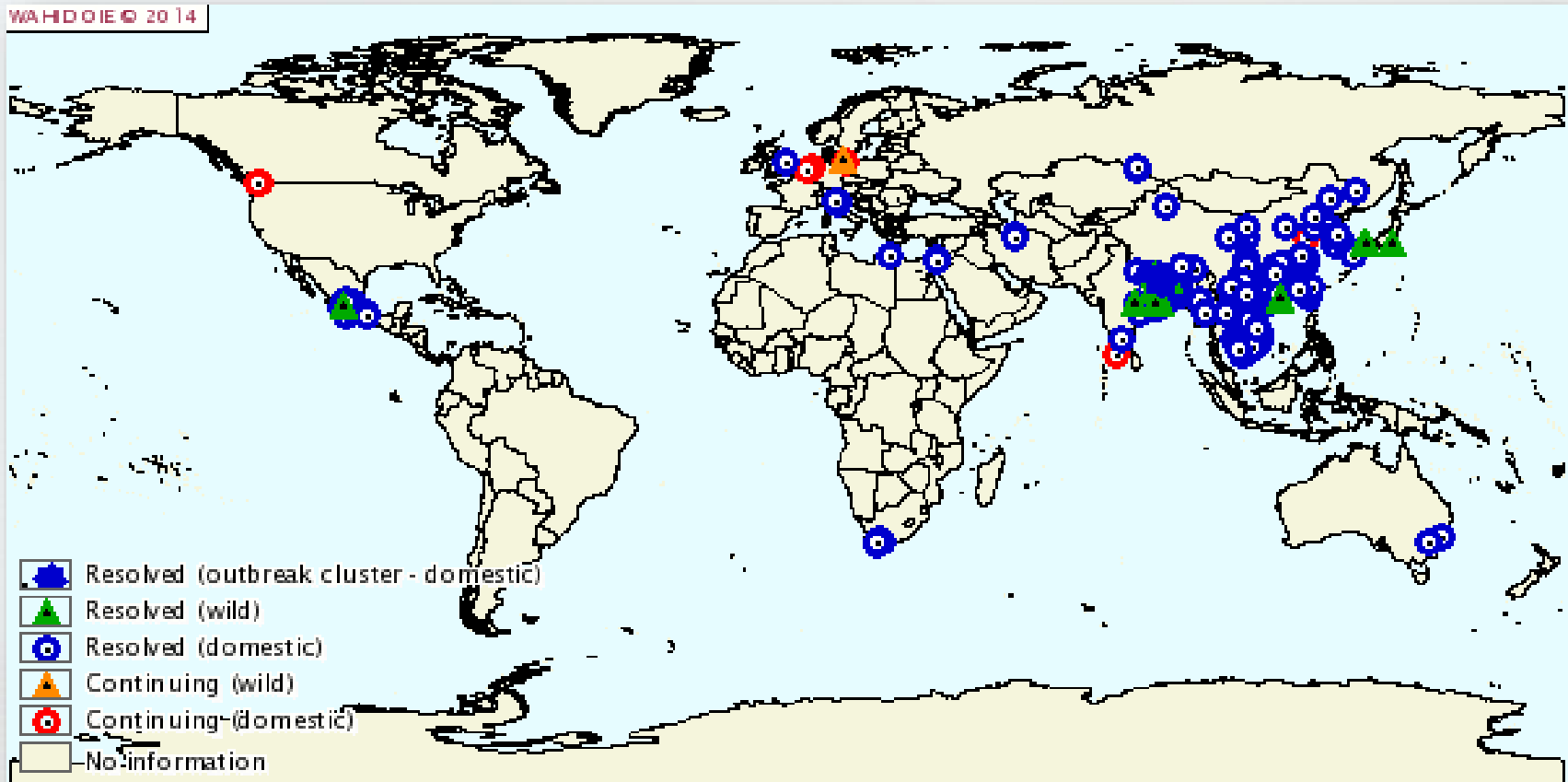
34. 2013: Italy, H7N7

35. 2013: Australia, H7N2

36. 2014: S. Korea and Europe, H5N8

37. 2014: Canada, H5N2

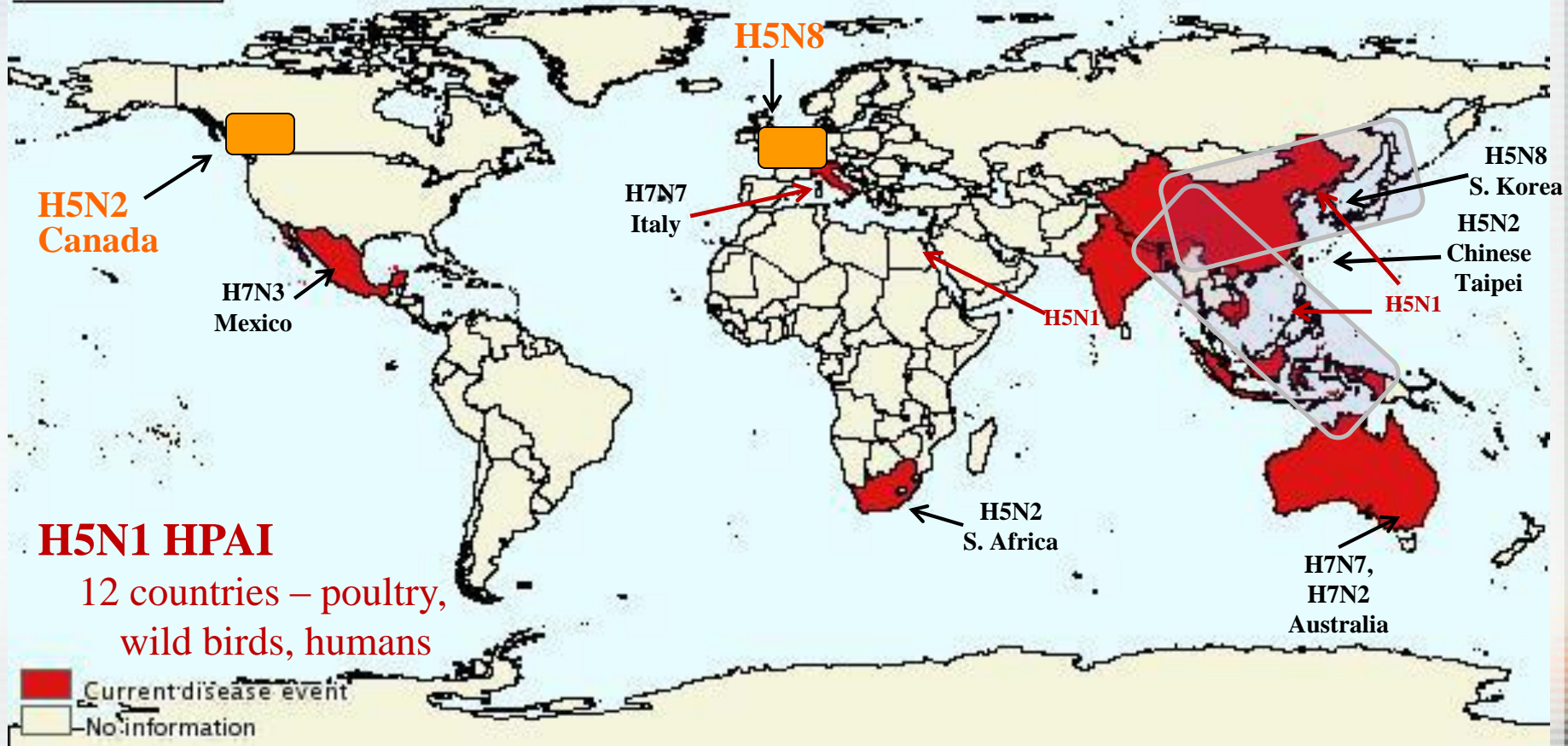
HPAI outbreaks reported to the OIE 2012-2014



H5 and H7 avian influenza in its low pathogenic form in poultry is a notifiable disease as per Chapter 10.4. on avian influenza of the Terrestrial Animal Health Code (2010)

HPAI (7/1/2012-2/7/2014): 18 countries

WAHID OIE © 2012



H5N2 HPAI

S. Africa – ostriches
Chinese Taipei – native chicken

H5N8 HPAI

S. Korea – ducks and other poultry

H7N2 HPAI

Australia - layers

H7N3 HPAI

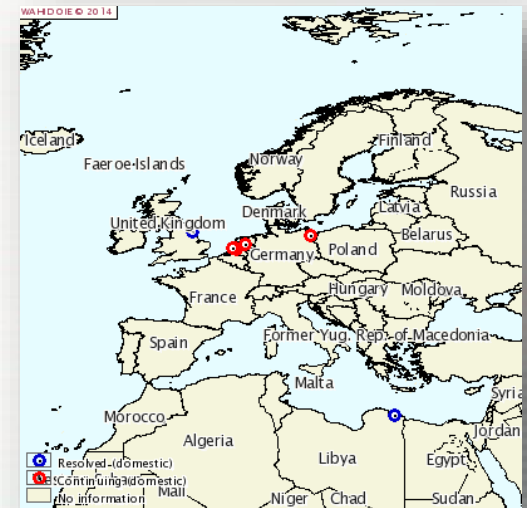
Mexico - layers

H7N7 HPAI

Italy – poultry
Australia - layers

H5N8 HPAI 2014

- Early this year, China, Japan and South Korea reported outbreaks of H5N8 in poultry farms as well as findings in migratory birds
- South Korea: 26+ outbreaks: breeding ducks, meat ducks, layer chickens, broilers, breeders. 15,114 deaths and 494,346 culled
- Since early November 2014, H5N8 HPAI was reported at poultry farms in Germany (1 farm), Netherlands (3 farms), and the UK (1 farm)
- A sample collected from an wild Common Teal legally hunted in Germany tested positive for HPAI H5N8



H7N3 HPAI in Mexico

- Reported to OIE 6/21/2012 – Jalisco
 - 44 farms
 - 10.6 million chickens affected– all layers
 - 11 million culled in outbreak zone
- Temporary vaccination authorized: 165.9 m doses used by Jan/2013
- Limiting factors in control:
 - Large number of diagnostic testing: 730 premises in Jalisco
 - Labor intensive depopulation of individual layers
- Resurgence in 2013 (central Mexico):
 - 12/1/2013 to 31/8/2013 – 64 outbreaks in Jalisco, Aguascalientes, Guanajuato and Puebla
 - 550,322 deaths, 6,230,022 culled
 - Layers, broiler breeders, backyard poultry and broilers
- H7N3 total – 110 outbreaks, 20 million poultry



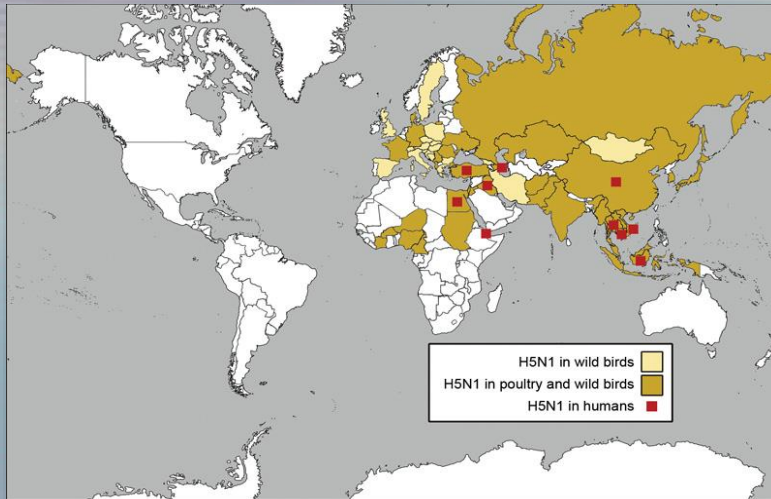
H5N2 HPAI in Canada

- Canadian health officials confirmed H5N2 HPAI in two of the four H5-related poultry outbreaks in British Columbia's Fraser Valley near Vancouver



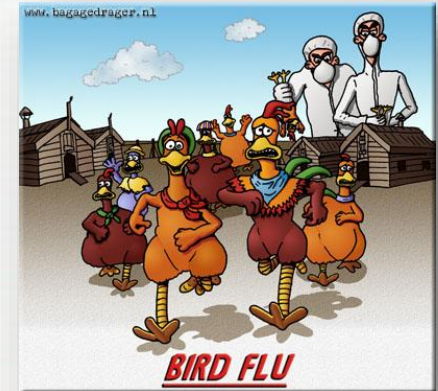
Up to 140,000 chickens and turkeys culled in B.C. as officials try to contain outbreak of highly-contagious avian flu

[T. Hopper | December 7, 2014](#). The National Post



Asian lineage H5N1 HPAI

H5N1 HPAI –The bird Flu

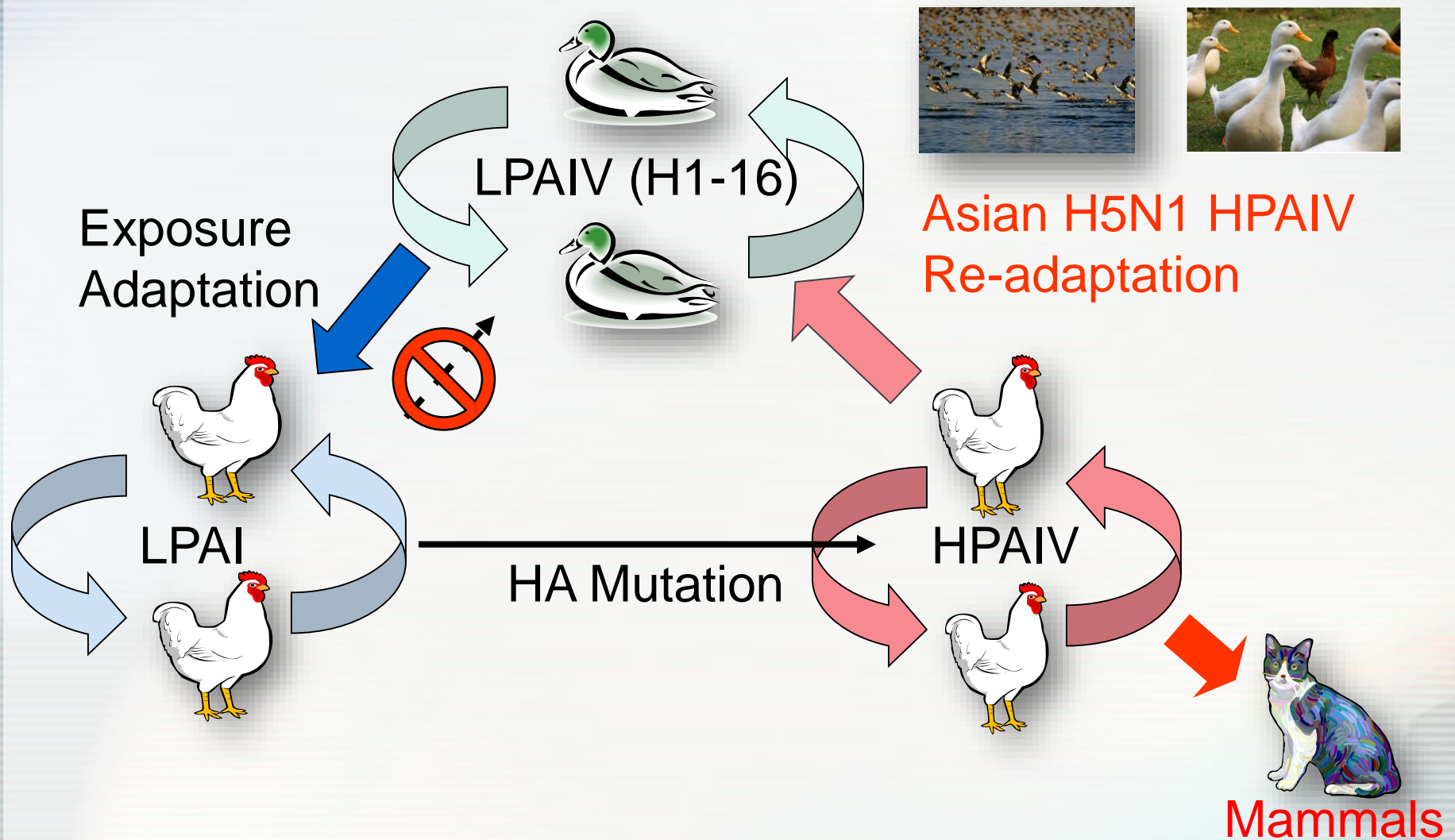


H5N1 HPAI

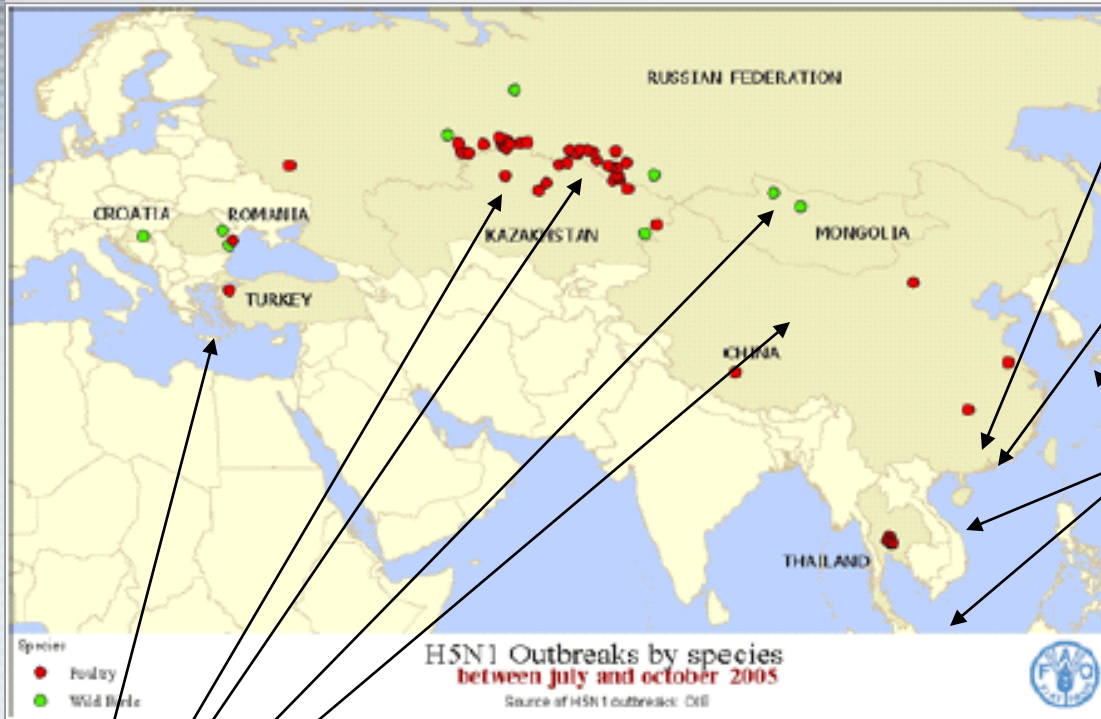
- H5N1 HPAI viruses continue to circulate in poultry and cause disease, and remain a threat to human and animal health (OIE-FAO-WHO)
- Outbreaks in poultry have seriously impacted livelihoods, the economy and international trade in affected countries



Ecology and Epidemiology - H5N1 HPAI



H5N1 HPAI Epizootics



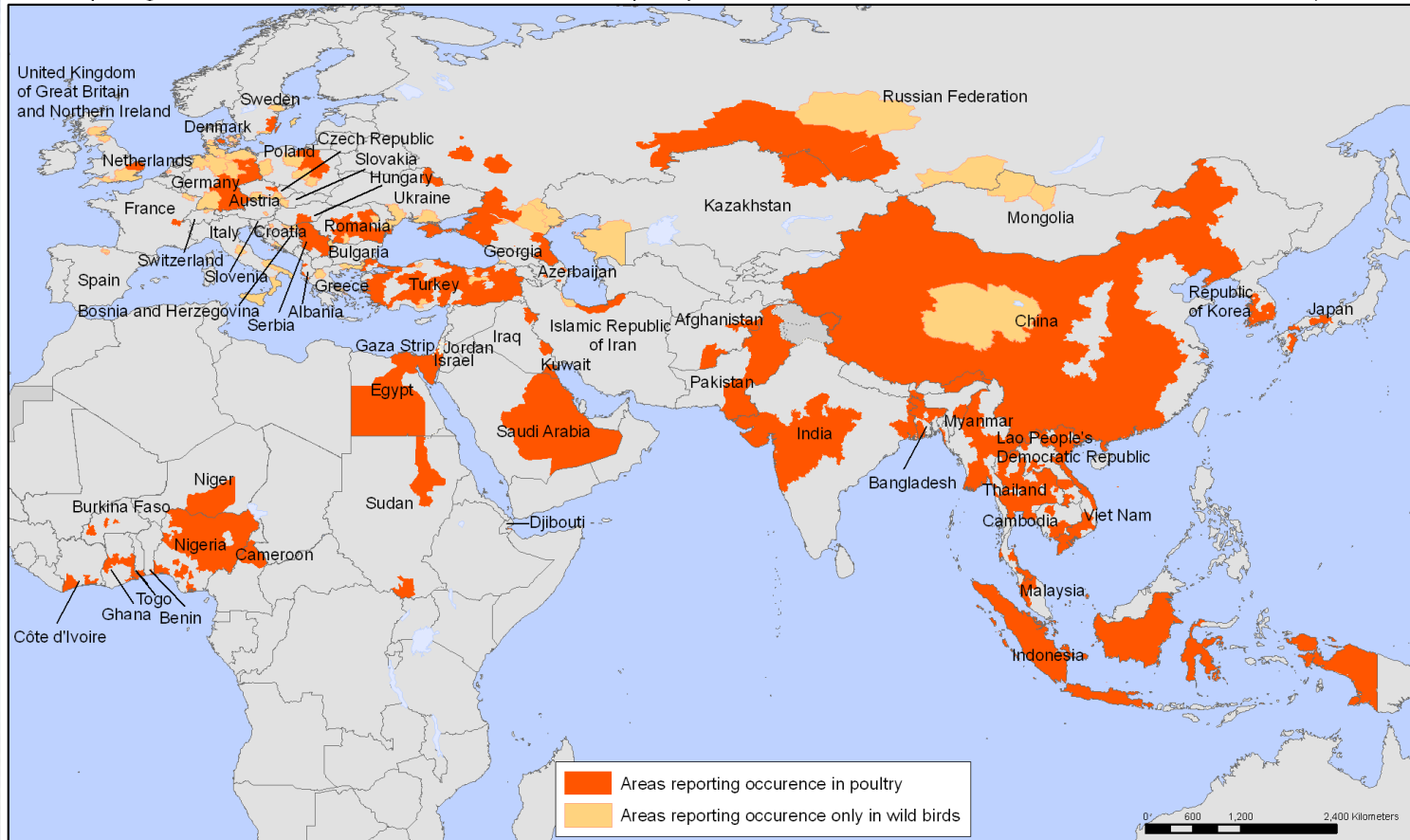
4. **Central Asia & E. Europe, wild bird –**
 - **Mid-2005: China (Qinghai Lake), Russia, Mongolia, Kazakhstan**
 - **Late-2005: Turkey, Romania, Kuwait, Croatia, Ukraine, Cyprus**

1. **Beginning - China: 1996-2006**
2. **Local extension - Hong Kong: 1997, 2001-3**
3. **SE Asian Regional Extension**
 - **S. Korea: 2003-4**
 - **Vietnam: 2004-6**
 - **Japan: 2004**
 - **Thailand: 2004-6**
 - **Cambodia: 2004-6**
 - **Laos: 2004-6**
 - **Taiwan (smuggled ducks): 2003**
 - **Indonesia: 2003-6**
 - **Malaysia: 2004**

H5N1 HPAI: 2006-2008

Areas reporting confirmed occurrence of H5N1 avian influenza in poultry and wild birds since 2003

Status as of 03 March 2008
Latest available update



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The boundaries and names shown and the designations used on this map do not imply the expression of any opinion whatsoever on the part of the World Health Organization concerning the legal status of any country, territory, city or area or of its authorities, or concerning the delimitation of its frontiers or boundaries. Dotted lines on maps represent approximate border lines for which there may not yet be full agreement.

Data Source: World Organisation for Animal Health (OIE)
and national governments
Map Production: Public Health Mapping and GIS
World Health Organization

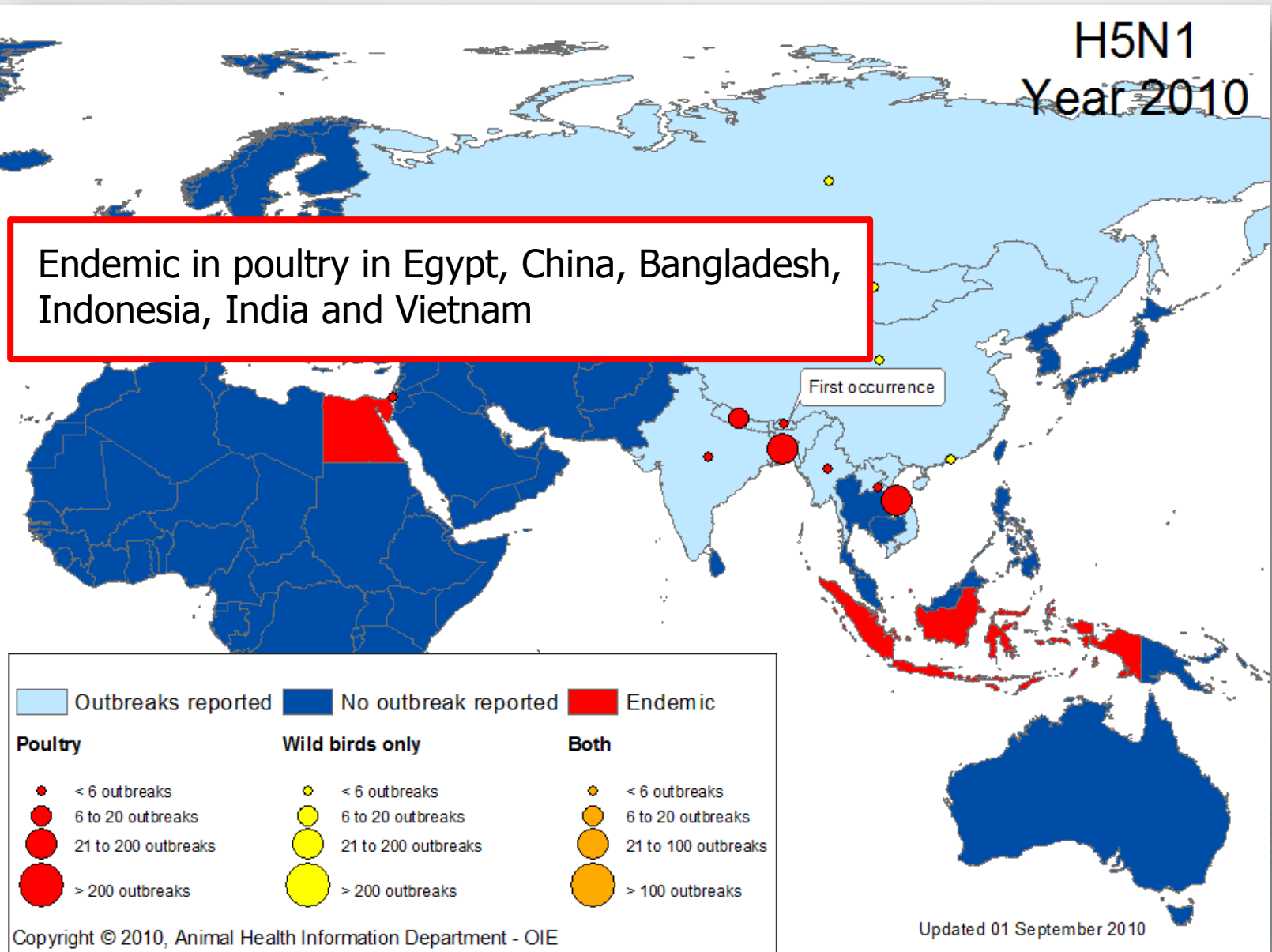
Africa
India
Bangladesh
Malaysia
Russia
Myanmar
Israel
Laos
Poland
etc.

- **+55 countries with cases in wild birds and/or poultry**
- **Over 220 million birds dead or culled 2004-2008 (FAO)**

H5N1 HPAI

H5N1
Year 2010

Endemic in poultry in Egypt, China, Bangladesh, Indonesia, India and Vietnam

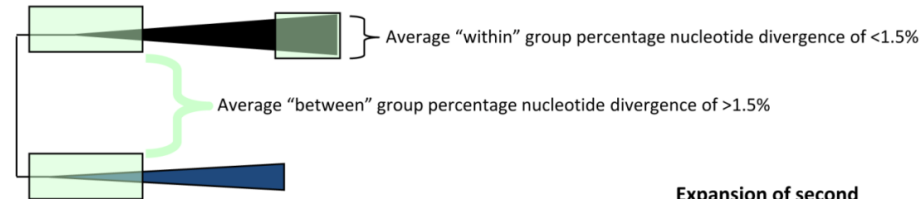


H5N1 HPAI in Vietnam

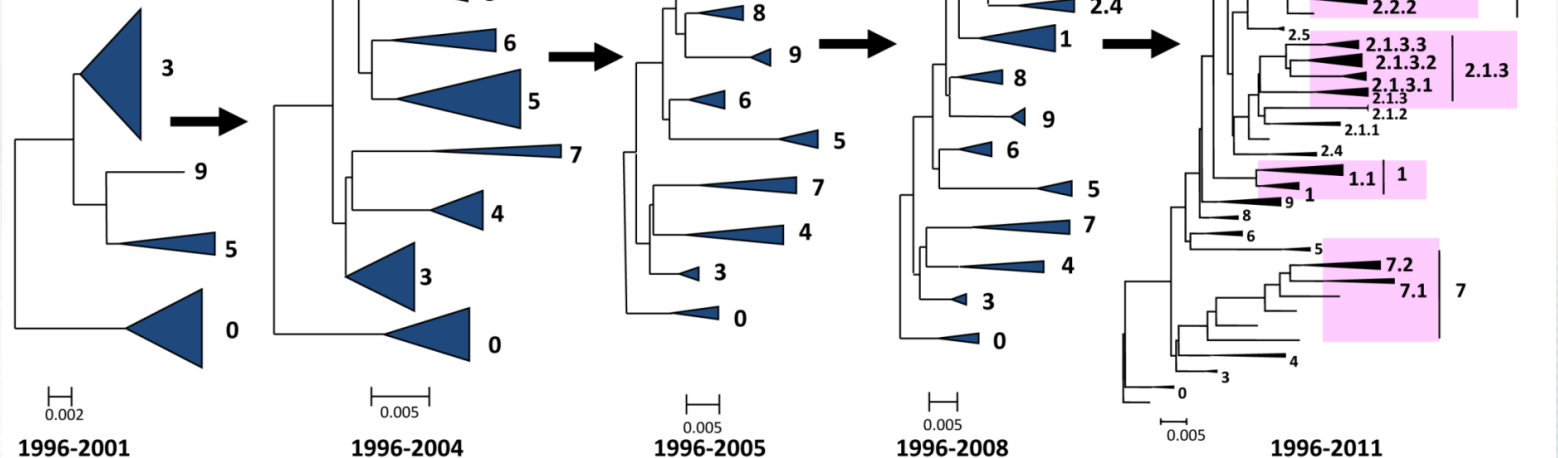


H5N1 HPAI viruses continue evolving

Evolution of the Asian H5 Hemagglutinin

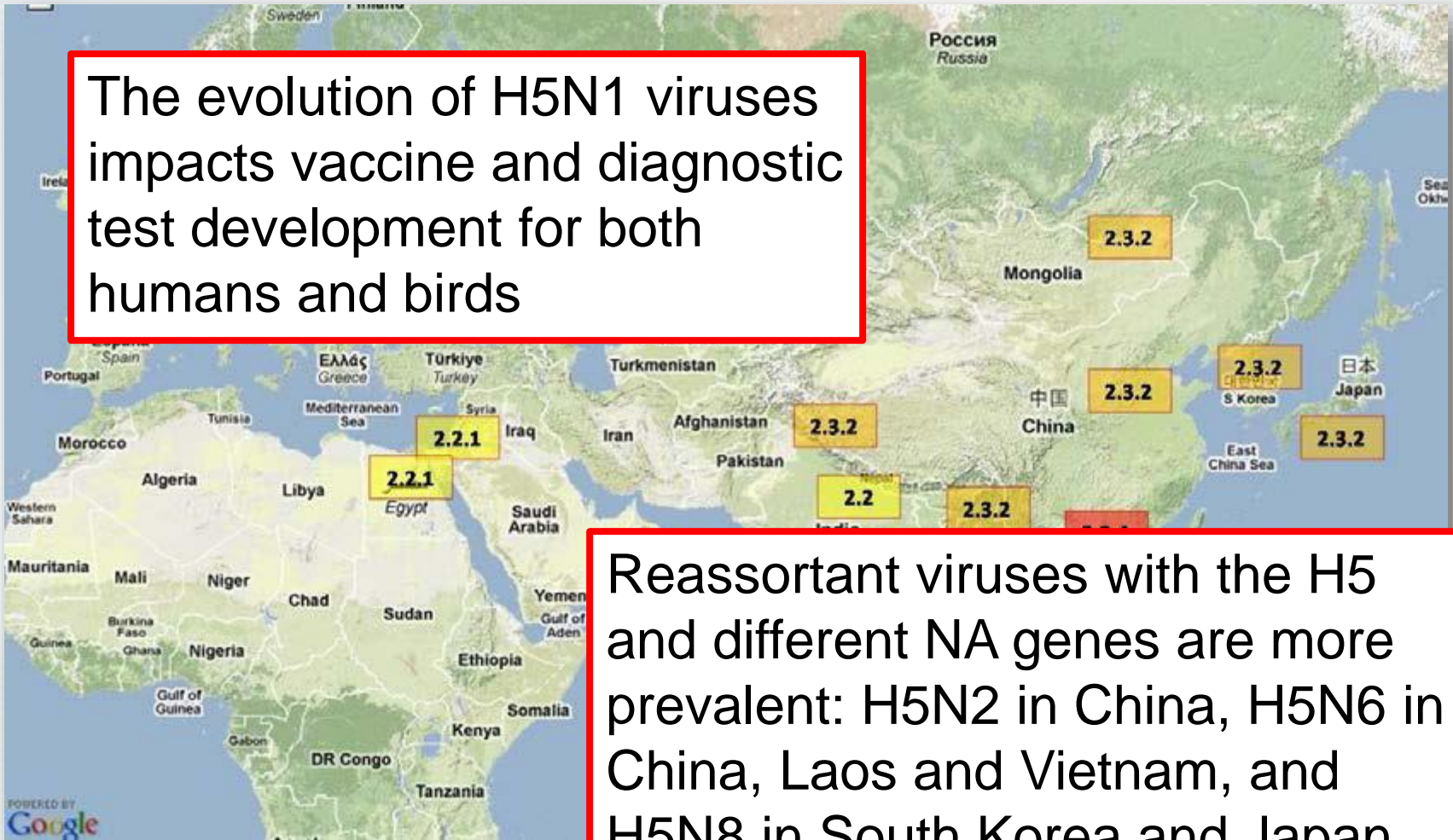


When discrete monophyletic groups begin to appear within a specific clade and those groups meet the nucleotide divergence criteria (as well as having bootstrap values >60), they are split into second order clades (but still considered part of the original first order clade). As a second order clade continues to evolve it may reach a similar level of genetic diversity at which point it may be split into third order clades and so on. The same clade designation criteria apply to first, second, and any higher order clade designations.



H5N1 HPAI viruses continue evolving

The evolution of H5N1 viruses impacts vaccine and diagnostic test development for both humans and birds



Reassortant viruses with the H5 and different NA genes are more prevalent: H5N2 in China, H5N6 in China, Laos and Vietnam, and H5N8 in South Korea and Japan

H5N1 HPAI and Public Health

Country	Total 2003-2014	
	cases	deaths
Azerbaijan	8	5
Bangladesh	7	1
Cambodia	56	37
China	47	30
Djibouti	1	0
Egypt	177	63
Indonesia	197	165
Iraq	3	2
Lao	2	2
Myanmar	1	0
Nigeria	1	1
Pakistan	3	1
Thailand	25	17
Turkey	12	4
Viet Nam	127	64
Total	668	393

Cumulative Number of Confirmed Human Cases of H5N1. Reported to WHO. 2003-2014



H5N1 HPAI and Public Health

- Human cases of H5N1 infection are rare and sporadic events, occurring mostly in areas where the virus is circulating endemically in poultry
- Exposure risks for human infection
 - Exposure one week before illness to poultry, especially direct handling of sick or dead poultry
 - Women- preparers of food
 - Limited human-to-human transmission
- H5N1 HPAI has not been a Food Safety issue
 - Virus is killed by pasteurization and cooking

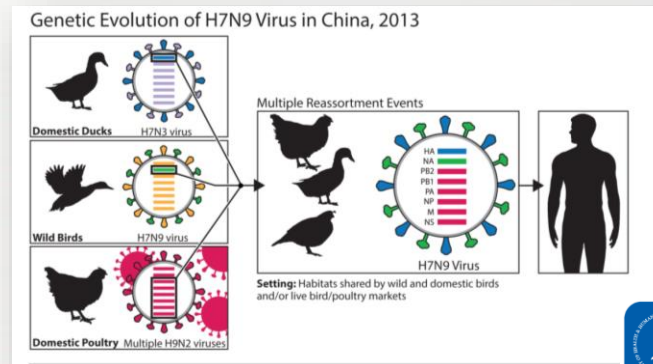


Natural Human AI Virus Infections

Year	Country	Subtype	Cases	Deaths
1959	USA	H7N7 HPAI	1	0
1978-9	USA	H7N7 LPAI	?	0
1996	U. Kingdom	H7N7 LPAI	1	0
1999-2011	China-Hong Kong	H9N2 LPAI	7+	0
2002-3	USA	H7N2 LPAI	2	0
2003	Netherlands	H7N7 HPAI	89	1
2004	Canada	H7N3 LP/HPAI	2	0
2006	U. Kingdom	H7N3 LPAI	1	0
2007	U. Kingdom	H7N2 LPAI	4	0
2012	Mexico	H7N3 HPAI	2	0
2003-2014	Asia/ME/Africa	H5N1 HPAI	650	389
2013-2014	China	H7N9 LPAI	463	127
2013-2014	China	H10N8 LPAI	3	2

H7N9 influenza in China

- A novel H7N9 influenza A virus emerged in humans eastern China in February of 2013. Cases were often severe and the case fatality rate approximately 32%
- The viral genome was rapidly characterized and was found to be a novel combination of genes from avian influenza viruses



- An avian reservoir was immediately suspected as a likely origin of human infections because of the genes of the virus were of avian origin and because recent contact with poultry was documented with numerous human cases

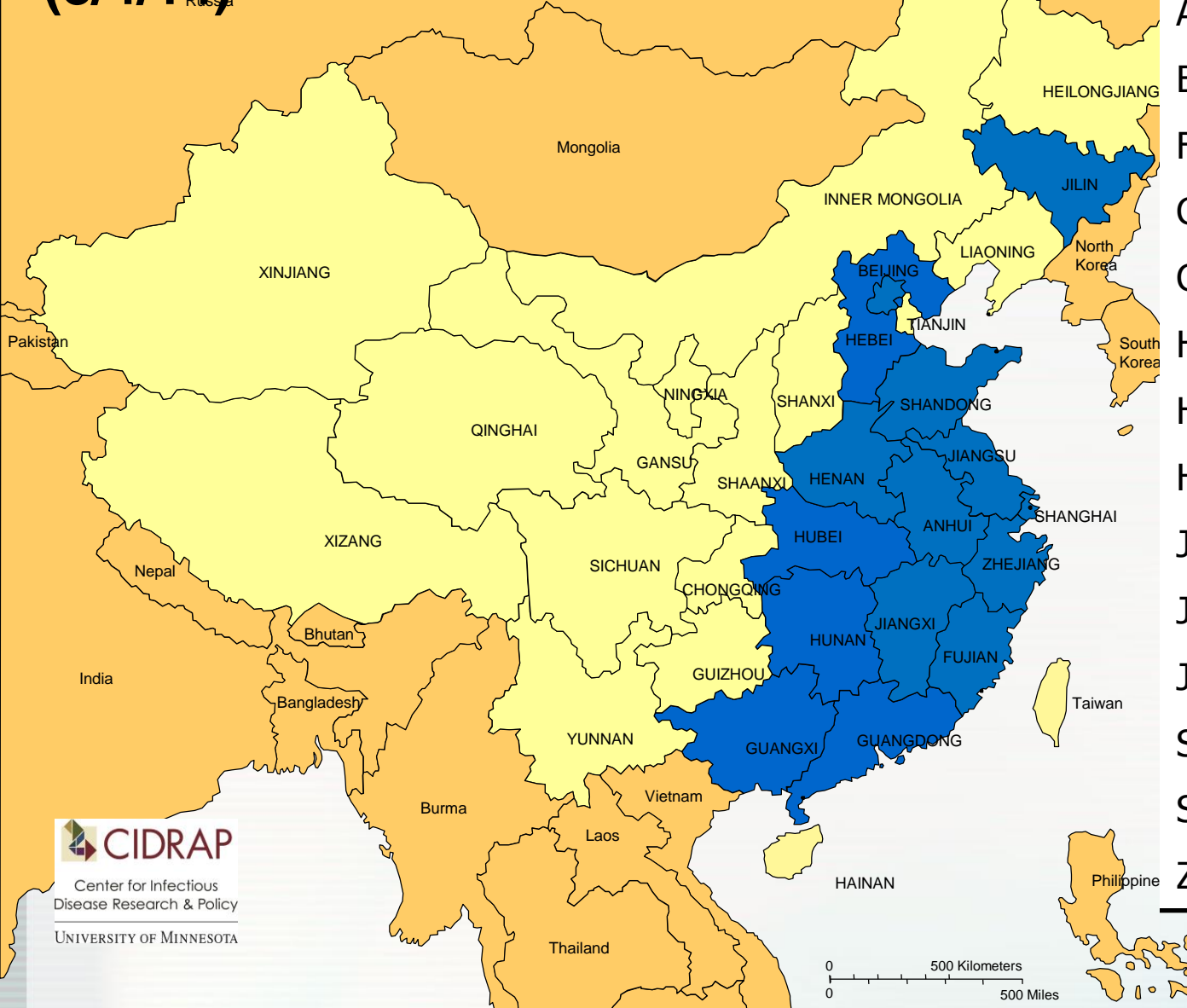
H7N9 influenza in China

- Chinese veterinary officials quickly started testing poultry associated with live poultry markets (LPM's), commercial poultry operations, and wild birds in the regions where human infections were being reported
- H7N9 viruses were detected in avian species in the LPM's including chickens, pigeons, ducks, and the environment
- Therefore LPM's were suspected of being a major source of human infections, and Chinese officials required closure of LPM's resulting in the reduction of human cases



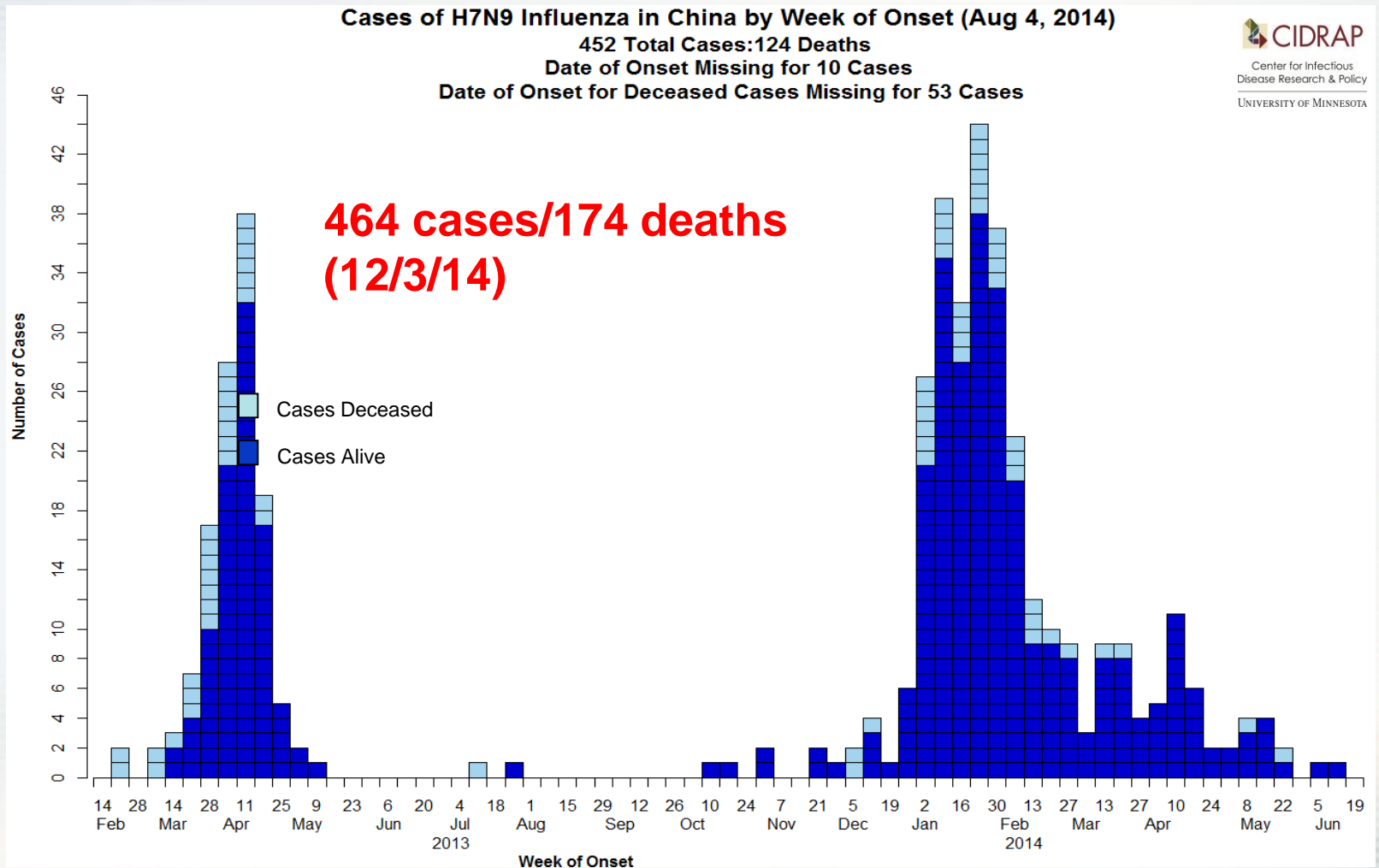
It's a LPAI

Location of H7N9 Influenza in China (8/4/14)*



Province/ City	Number of Cases
Anhui	18
Beijing	5
Fujian	22
Guangdong	118
Guangxi	3
Hebei	1
Henan	4
Hunan	24
Jiangsu	61
Jiangxi	8
Jilin	2
Shandong	4
Shanghai	42
Zhejiang	140

H7N9 influenza human cases



Since 2 December 3 new human cases have been reported in **Guangdong** and **Fujian Provinces** and **Shanghai City**

<http://empres-i.fao.org/eipws3g/>

Role of LPM's

- Retail LPMs provide a mixing ground for the emergence of novel strains of influenza
 - The closure of 780 LPM's in the Chinese cities of Shanghai, Hangzhou, Huzhou and Nanjing in April 2013 reduced the daily number of H7N9 infections by more than 97 percent (*Cowling et al, The Lancet 2014*)
 - Losses associated with the closures have been estimated at about 57 billion yuan (about \$9 billion/ 7 billion euros)
- The findings confirm that LPM closure, though a huge economic setback, is a highly effective intervention to prevent human disease and protect public health



Avian Influenza: Prevention and Control

Avian Influenza

Animal Health Risk and Impact



- Economics: Treatment costs and production impacts
- Animal suffering and welfare
- Negative impact on human nutrition and livelihoods (i.e. Food Security)
- Trade impact: real-risk and pseudo-risk with outcome of trade barriers

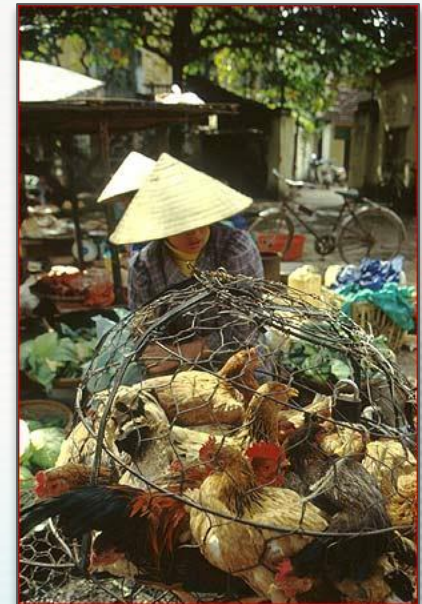
Avian Influenza

Economic losses from AI result from:

- Direct losses:
 - Depopulation and disposal
 - High morbidity and mortality
 - Quarantine and surveillance
 - Indemnities
- Indirect losses:
 - loss of consumer confidence in poultry products
 - costs to prevent, manage or eradicate the disease
 - effect on trade
- HPAI negatively impacts the livelihoods of millions of people especially the rural poor
- Early and successful control of AI requires an accurate and rapid diagnosis



<http://www.medindia.net/news>



Control of Avian Influenza

- In most countries, LPAI and HPAI are not common so prevention is the primary goal
- **Surveillance and diagnostics**
- When AI occurs, eradication is the overall objective
 - Quarantine, depopulation, cleaning and disinfection
- **Appropriate biosecurity**
 - Control human traffic
 - Introduction of new birds into flock
 - Avoid open range rearing in waterfowl prevalent areas
- Vaccination can be used as part of a control program during an outbreak
- Wide variation among countries in regards to veterinary diagnostics and animal health infrastructure



In the USA

- HPAI is rare
 - 1924-25: Classic Fowl Plague (H7N7?), New England and upper Midwest states
 - 1929: Classic Fowl Plague (H7N7?), New Jersey
 - 1983-84: H5N2, Pennsylvania, Virginia and Maryland
 - 2004: H5N2, Texas
- H5/H7 LPAI – historically handled by companies and states; since 2005, national control program (prevent H5/7 LPNAIV → HPAIV)
- Other LPAI (H1-4, H6, H8-16) – no federal program but handled by companies and states

Summary

- Avian Influenza viruses are constantly emerging and changing, and will always be a threat to animal and human health
- Both animal and public health sectors at the national, regional and international levels should maintain vigilance in regularly detecting, reporting, and characterizing animal influenza viruses, and in assessing and managing existing and evolving health risks associated with these viruses

Thanks for your attention!

